

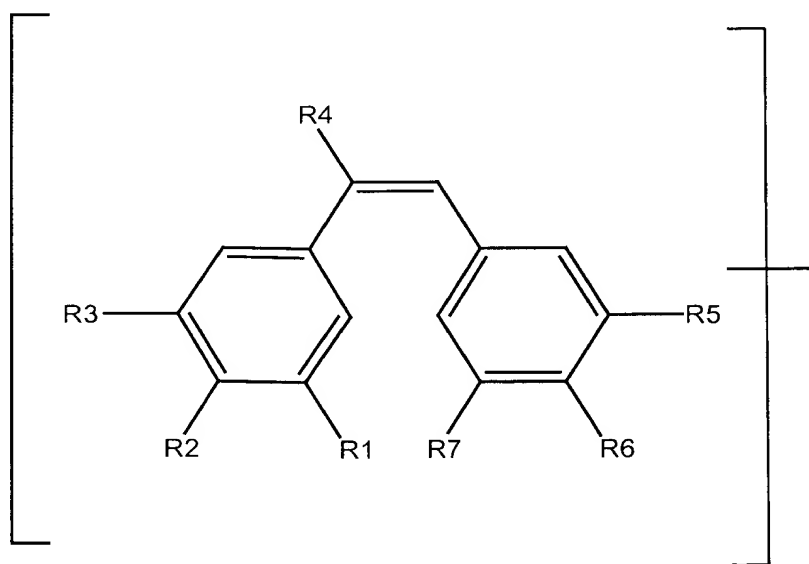
Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims

Claims 1-20 (Cancelled)

21 (Previously submitted). A compound of formula AXB useful in inducing necrosis in vascular tissue of a tumor in a mammal, said compound containing (a) a first moiety, A, which is a cis-stilbene moiety of formula II



wherein R1, R2 and R3 are each independently H, optionally substituted alkoxy, optionally substituted alkyl or halogen

R4 is hydrogen or cyano

R5, R6 and R7 are each independently H, hydroxy, optionally substituted alkyl, halogen, amino, alkylamino, dialkylamino, cyano, nitro, carboxyl, alkanoyl, alkoxy carbonyl, alkoxy carbonylamino, aminocarbonylamino, alkylaminocarbonylamino, dialkylaminocarbonylamino, alkylcarbonylamino, alkylsulphonyl, aminosulphonyl, alkylaminosulphonyl, dialkylaminosulphonyl, alkylsulphonylamino, aminosulphonylamino, alkylaminosulphonylamino, dialkylaminosulphonylamino, mercapto, alkylsulphanyl, or alkylsulphanyl,

with the proviso that at least ~~to~~ two of R1, R2 and R3 must be optionally substituted alkoxy, and (b) a second moiety, B, which is an inhibitor of nitric oxide synthase said first and second moieties being coupled in the compound by a linker bond, atom or group X bound to any available valency of A such that the compound has an increased activity in inducing necrosis in said vascular tissue as compared with a compound containing said first moiety without the second moiety

or a hydrate or pharmaceutically acceptable salt of such a compound.

22 (Canceled)

23 (Previously submitted). The compound according to claim 21, wherein the compound is a hydrate, or a pharmaceutically acceptable salt thereof.

24 (cancelled). .

25 (currently amended) The compound according to claim ~~24~~ 21, in which ~~the first and second moieties are coupled through a linker group~~ X is selected from the group consisting of an optionally substituted methylene chain, and $-(CH_2)_m-Y-(CH_2)_n-$ wherein Y is selected from -O-, -S-, SO_2 -, NH-, Nalkyl-, -CO-, -OC(O)-, -NHC(O)-, -N(alkyl)C(O)-, -NHC(O)NH-, NalkylC(O)NH-, NalkylC(O)Nalkyl-, -NHSO₂-, NalkylSO₂-, NHSO₂NH-, NalkylSO₂NH-, NalkylSO₂Nalkyl- and -OC(O)O-, m is 0-3 and n is 0-3.

26 (Previously submitted) The compound according to claim 21, in which the second moiety is selected from the group consisting of an amino acid inhibitor of nitric oxide synthase, a thiocitrulline derivative, an S-alkylisothiourea derivative and a 2-aminopyridine derivative.

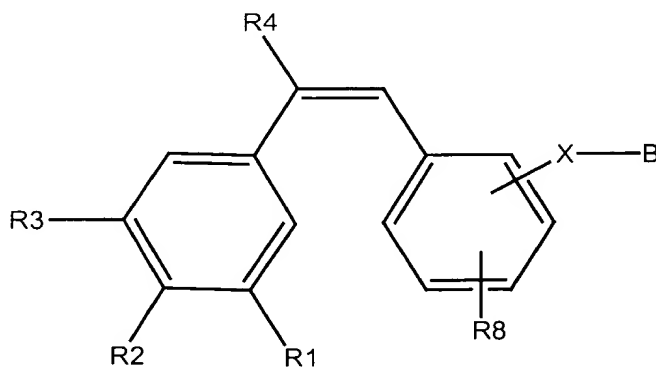
Claim 27 (Previously submitted) The compound according to claim ~~22~~ 21, wherein the second moiety is a group $-C(O)CH(NH_2)-CH_2p-NHC(NH)Z$ wherein p is 1-5 and Z is alkyl, alkylamino, dialkylamino, nitroamino, hydrazino or alkylthio, or a group $-NHCH(CO_2R_{10})-(CH_2)p-NHC(NH)Z$ and R_{10} is hydrogen or alkyl.

Claim 28 (Previously submitted) The compound according to claim ~~22~~ 21, wherein the second moiety is a group $-C(O)CH(NH_2)-CH_2p-NHC(S)NH_2$ or a group $-NHCH(CO_2R_{10})-(CH_2)p-NHC(S)NH_2$.

Claim 29 (Previously submitted) The compound according to claim ~~22~~ 21, wherein the second moiety is $-(CH_2)p-SC(NH)NH_2$.

Claim 30 (Previously submitted) The compound according to claim ~~22~~ 21, wherein the second moiety is 4-methyl-2-pyridinylamino.

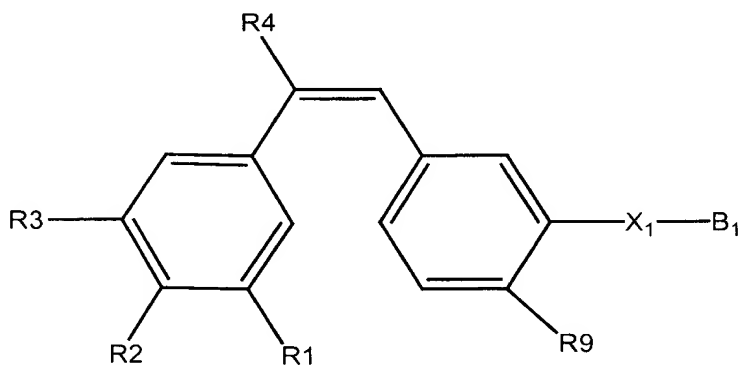
Claim 31 (Previously submitted) The compound according to claim 21, wherein the compound is



wherein B is the second moiety; and X is a linker bond, atom or group as defined in claim 21; and R8 is alkyl, amino, hydroxy, alkoxy or halogen.

Claim 32 (Previously submitted) The compound according to claim 31, wherein X is -O- or -NH- and B is a group $-C(O)CH(NH_2)-(CH_2)_p-NHC(NH)Z$, wherein p is 1-5 and Z is alkyl, alkylamino, dialkylamino, nitroamino, hydrazino or alkylthio or a group $-NHCH(R_{10})-CH_2)_p-NHC(NH)Z$ and wherein R₁₀ is hydrogen or alkyl.

Claim 33 (Currently amended) The compound according to claim 32, wherein the compound is



wherein

R₉ is alkyl, alkoxy or halogen

X₁ is O or NH

B₁ is a group $-C(O)CH(NH_2)_p-NHC(NH)Z$ wherein p is 1-5 and Z is alkyl, alkylamino, dialkylamino, nitroamino, hydrazino or alkylthio.

34 (Previously submitted) The compound according to claim 21, wherein the compound is selected

from the group consisting of

(Z)-1-(4-methoxy-3-N^G-nitroarginyloxyphenyl)-2-(3,4,5-trimethoxyphenyl)ethene

(Z)-N-[2-methoxy-5-[2-(3,4,5-trimethoxyphenyl)ethenyl]phenoxy carbonyl]N^G- nitroarginine methyl ester;

(Z)-N-[2-methoxy-5-[2-(3,4,5-trimethoxyphenyl)ethenyl]phenoxy carbonyl]N^G- nitroarginine; and

(Z)-N-[2-methyl-5-[2-(3,4,5-trimethoxyphenyl)ethenyl]phenoxy carbonyl]N^G nitroarginine methyl ester.

35 (Previously submitted) The compound according to claim 21, wherein the first and second moieties are coupled through a linker bond.

36 (currently amended). A method for inducing necrosis in vasculature of a tumor in a mammal, comprising administering to the ~~animal~~ mammal the compound of claim 34 in an amount effective for said inducing.

37 (currently amended). A method for inducing necrosis in vasculature of a tumor in a mammal, comprising administering to the ~~animal~~ mammal the compound of claim 21 in an amount effective for said inducing.

38 (currently amended). A method for inducing necrosis in vasculature of a tumor in an animal, comprising administering to the ~~animal~~ mammal the compound of claim 24 in an amount effective for said inducing.

39 (Previously submitted). A method for inducing necrosis in vasculature of a tumor in a mammal, comprising administering to the mammal the compound of claim 27 in an amount effective for said inducing.

40 (currently amended). A method for inducing necrosis in vasculature of a tumor in a mammal ~~animal~~, comprising administering to the animal the compound of claim 31 in an amount effective for said inducing.